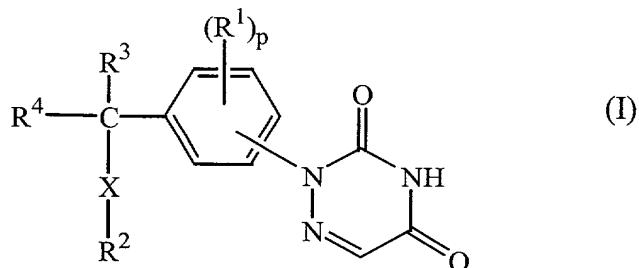


CLAIMS

1. A compound having the formula:



5      the N-oxides, the pharmaceutically acceptable addition salts and the stereochemically isomeric forms thereof, wherein :

p represents an integer being 0, 1, 2, 3 or 4;

X represents O, S, NR<sup>5</sup> or a direct bond or-X-R<sup>2</sup> taken together may represent cyano;

10     Y represents O, S, NR<sup>5</sup>, or S(O)<sub>2</sub>;

each R<sup>1</sup> independently represents C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkyl, halo, polyhaloC<sub>1-6</sub>alkyl, hydroxy, mercapto, C<sub>1-6</sub>alkyloxy, C<sub>1-6</sub>alkylthio, C<sub>1-6</sub>alkylcarbonyloxy, aryl, cyano, nitro, Het<sup>3</sup>, R<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup> or C<sub>1-4</sub>alkyl substituted with C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, R<sup>6</sup> or NR<sup>7</sup>R<sup>8</sup>;

15     R<sup>2</sup> represents Het<sup>1</sup>, C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkyl substituted with one or two substituents selected from C(=O)-Z-R<sup>14</sup>, hydroxy, mercapto, cyano, amino, mono- or di(C<sub>1-4</sub>alkyl)amino, C<sub>1-6</sub>alkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkylthio optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkylsulfonyloxy, C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, aryl, aryloxy, arylthio, Het<sup>1</sup>, Het<sup>1</sup>oxy and Het<sup>1</sup>thio; and if X is O, S or NR<sup>5</sup>, then R<sup>2</sup> may also represent aminothiocarbonyl, C<sub>1-4</sub>alkylcarbonyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-4</sub>alkylthiocarbonyl optionally substituted with C(=O)-Z-R<sup>14</sup>, arylcarbonyl, arylthiocarbonyl, Het<sup>1</sup>carbonyl or Het<sup>1</sup>thiocarbonyl;

20     R<sup>3</sup> represents hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl;

R<sup>4</sup> represents hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl; or

R<sup>3</sup> and R<sup>4</sup> taken together form a C<sub>2-6</sub>alkanediyl;

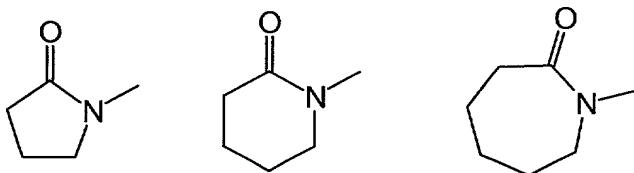
R<sup>5</sup> represents hydrogen or C<sub>1-4</sub>alkyl;

each R<sup>6</sup> independently represents C<sub>1-6</sub>alkylsulfonyl, aminosulfonyl,

piperidinylsulfonyl, mono- or di(C<sub>1-4</sub>alkyl)aminosulfonyl, mono- or

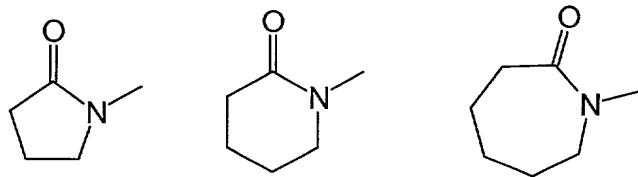
di(benzyl)aminosulfonyl, polyhaloC<sub>1-6</sub>alkylsulfonyl, C<sub>1-6</sub>alkylsulfinyl,

5 phenylC<sub>1-4</sub>alkylsulfonyl, piperazinylsulfonyl, aminopiperidinylsulfonyl,  
piperidinyl-aminosulfonyl, N-C<sub>1-4</sub>alkyl-N-piperidinylaminosulfonyl, Y-R<sup>14</sup>, mono-  
or di-(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkylsulfonyl, Het<sup>6</sup>sulfonyl or C<sub>3-7</sub> cycloalkylsulfonyl;  
each R<sup>7</sup> and each R<sup>8</sup> are independently selected from hydrogen, C<sub>1-4</sub>alkyl,  
hydroxyC<sub>1-4</sub>alkyl, mercapto-C<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, aryl, arylC<sub>1-4</sub>alkyl,  
10 C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, C<sub>1-4</sub>alkyl-thiocarbonyl, arylcarbonyl,  
aryltiocarbonyl, Het<sup>3</sup>thiocarbonyl, Het<sup>3</sup>carbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl,  
arylaminocarbonyl, arylaminothiocarbonyl, Het<sup>3</sup>aminocarbonyl,  
Het<sup>3</sup>aminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl,  
C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>,  
15 Het<sup>4</sup> and R<sup>6</sup>; or R<sup>7</sup> and R<sup>8</sup> taken together with the nitrogen atom to which they  
are attached form a radical of formula



R<sup>9</sup> and R<sup>10</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl,  
hydroxyC<sub>1-4</sub>alkyl, mercapto-C<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl,

20 phenylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, arylcarbonyl,  
Het<sup>3</sup>carbonyl, Het<sup>3</sup>thiocarbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl,  
arylaminocarbonyl, arylaminothiocarbonyl, Het<sup>3</sup>aminocarbonyl,  
Het<sup>3</sup>aminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl,  
C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>,  
25 Het<sup>4</sup> and R<sup>6</sup>; or R<sup>9</sup> and R<sup>10</sup> taken together with the nitrogen atom to which they  
are attached form a radical of formula

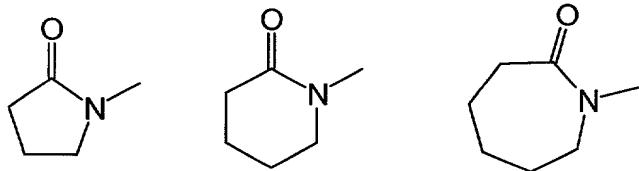


each R<sup>11</sup> independently being selected from hydroxy, mercapto, cyano, nitro,  
halo, trihalomethyl, C<sub>1-4</sub>alkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>,

C<sub>1-6</sub>alkylthio optionally substituted with C(=O)-Z-R<sup>14</sup>, formyl,

5 trihaloC<sub>1-4</sub>alkylsulfonyloxy, R<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup>, C(=O)NR<sup>15</sup>R<sup>16</sup>, -C(=O)-Z-R<sup>14</sup>,  
-Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, aryl, aryloxy, arylcarbonyl, arylthiocarbonyl,  
C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>3-7</sub>cycloalkyloxy  
optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>3-7</sub>cycloalkylthio optionally substituted  
with C(=O)-Z-R<sup>14</sup>, phthalimide-2-yl, Het<sup>3</sup>, Het<sup>4</sup>, C(=O)Het<sup>3</sup>, C(=O)C<sub>1-4</sub>alkyl

10 optionally be substituted with one or more substituents independently selected  
from hydroxy, mercapto, halo and phenyl;  
R<sup>12</sup> and R<sup>13</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl,  
hydroxyC<sub>1-4</sub>alkyl, mercapto-C<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl, phenyl-  
C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, C<sub>1-4</sub>alkylthiocarbonyl,  
15 arylcarbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, arylaminocarbonyl,  
arylaminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl,  
C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup> and  
R<sup>6</sup>; or R<sup>12</sup> and R<sup>13</sup> taken together with the nitrogen atom to which they are  
attatched form a radical of formula

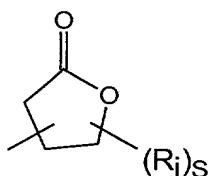


20 each R<sup>14</sup> independently represents hydrogen; C<sub>1-20</sub>acyl or C<sub>1-20</sub>alkylC<sub>1-20</sub>acyl  
(having a straight or branched, saturated or unsaturated hydrocarbon chain  
having 1 to 20 carbon atoms) optionally substituted with one or more  
substituents selected from hydroxy, mercapto, hydroxyC<sub>1-4</sub>alkyl, mercapto-  
C<sub>1-4</sub>alkyl, NR<sup>17</sup>R<sup>18</sup>, aryl, mono- or di-(C<sub>1-4</sub>alkyl)amino, cyano and Het<sup>5</sup>;

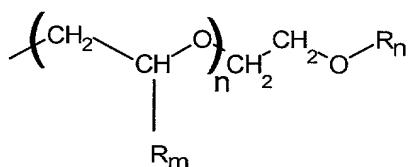
25

$C_{1-20}$ alkyl optionally substituted with one or more substituents selected from hydroxy, halo, mercapto,  $C_{1-4}$ alkyloxy $C_{1-4}$ alkyloxy, mercapto $C_{1-4}$ alkyl,  $NR^{17}R^{18}$ , aryl, mono- or di-( $C_{1-4}$ alkyl)amino, cyano,  $Het^5$ ,  $C_{1-4}$ alkyloxycarbonyl, aryl $C_{1-4}$ alkyloxycarbonyl, aryl $C_{1-4}$ alkyloxy, aryl $C_{1-4}$ alkylthiocarbonyl,

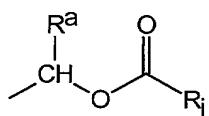
5    aryl $C_{1-4}$ alkylthio,  $Het^5C_{1-4}$ alkyloxy, aryl $C_{1-4}$ alkylthio,  $C_{3-7}$  cycloalkyl and  $Het^5C_{1-4}$ alkylthio;  $C_{3-20}$ alkenyl optionally substituted with phenyl;  $C_{3-20}$ alkynyl;  $C_{3-7}$  cycloalkyl optionally substituted with one or more substituents selected from hydroxy, mercapto, halo, mercapto $C_{1-4}$ alkyl and hydroxy $C_{1-4}$ alkyl;  $Het^5$  or phenyl or  $R^{14}$  represents a radical having any of the following formulae:



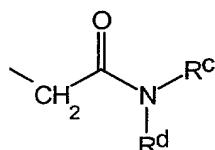
(a)



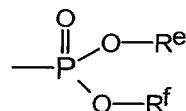
(b)



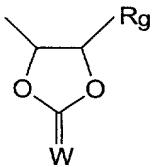
(c)



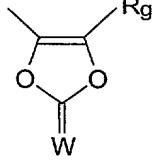
(d)



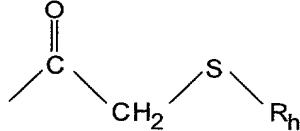
(e)



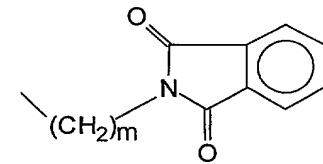
(h)



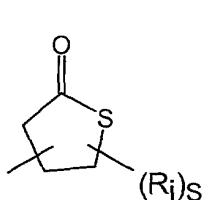
(i)



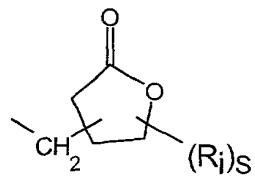
(j)



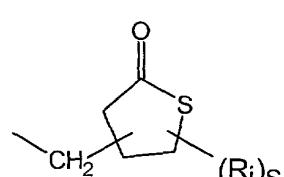
(k)



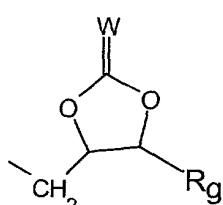
(1)



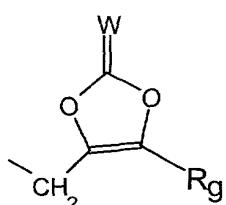
(m)



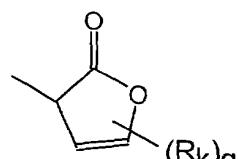
(n)



(0)



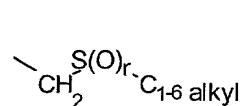
(p)



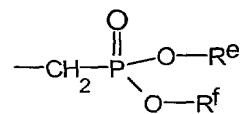
(q)



(r)



(S)



(t)

wherein m is 1 to 4, n is 0 to 5, q is 0 to 2, r is 0 to 2 and s is 0 to 4;

5 R<sup>b</sup> is selected from hydrogen, C<sub>1-6</sub>alkyl, phenyl, C<sub>3-7</sub>cycloalkyl, C<sub>1-4</sub> alkyloxyC<sub>1-6</sub>alkyl and C<sub>1-4</sub> alkyl-Y-C<sub>1-4</sub>alkyl;  
R<sup>a</sup>, R<sup>c</sup>, R<sup>d</sup>, R<sup>e</sup> and R<sup>f</sup> are each independently selected from hydrogen, C<sub>1-6</sub>alkyl, phenyl and C<sub>3-7</sub>cycloalkyl, or R<sup>e</sup> and R<sup>f</sup> taken together may form -CH<sub>2</sub>-CH<sub>2</sub>-, -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>- or -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>;-  
10 R<sub>g</sub>, R<sub>h</sub> and R<sub>k</sub> are each independently hydrogen or C<sub>1-4</sub> alkyl; R<sub>i</sub> is selected from hydroxy, C<sub>3-7</sub>cycloalkyl and C<sub>1-4</sub>alkyl, or two R<sub>i</sub> taken together may form -CH<sub>2</sub>-CH<sub>2</sub>-, -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>- or -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>- (thus building a spiro radical); R<sub>j</sub> is selected from -O-R<sub>b</sub>; C<sub>1-6</sub>alkyl optionally substituted with phenyl or

C<sub>3</sub>-7cycloalkyl; phenyl; C<sub>3</sub>-7cycloalkyl optionally substituted with C<sub>1</sub>-4 alkyloxy and mono- or di(C<sub>1</sub>-4alkyl)amino;

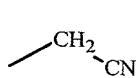
R<sub>m</sub> is hydrogen or C<sub>1</sub>-4 alkyloxy;

R<sub>n</sub> is hydrogen, C<sub>1</sub>-4alkyl, C<sub>3</sub>-7cycloalkyl, phenyl or phenylC<sub>1</sub>-4alkyl; and

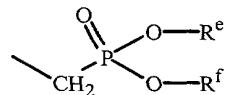
5 W represents O or S;

each Z independently represents O, S, NH, -CH<sub>2</sub>-O- or -CH<sub>2</sub>-S- whereby -CH<sub>2</sub>- is attached to the carbonyl group; or

-Z-R<sup>14</sup> taken together form a radical of formula



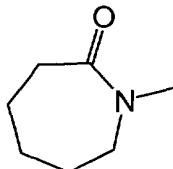
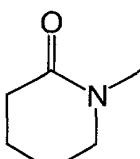
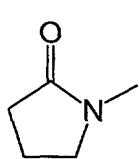
(f)



(g)

10 R<sup>15</sup> and R<sup>16</sup> are each independently selected from hydrogen; C<sub>1</sub>-4alkyl optionally substituted with one or more substituents independently selected from hydroxy, mercapto, aryl, mono- or di(C<sub>1</sub>-4alkyl) amino and pyridinyl; C<sub>1</sub>-4alkyloxy; aryl; -C(=O)-Z-R<sup>14</sup>; arylcarbonyl; arylthiocarbonyl; arylaminocarbonyl; arylaminothiocarbonyl; aminocarbonylmethylene; mono- or di(C<sub>1</sub>-4alkyl) aminocarbonylmethylene; Het<sup>3</sup>aminocarbonyl; Het<sup>3</sup>aminothio-carbonyl; pyridinylC<sub>1</sub>-4alkyl; Het<sup>3</sup> and R<sup>6</sup>; or R<sup>15</sup> and R<sup>16</sup> taken together with the nitrogen atom to which they are attached form a radical of formula

15



R<sup>17</sup> and R<sup>18</sup> are each independently selected from hydrogen, C<sub>1</sub>-6alkyl optionally substituted with one or more substituents independently selected from hydroxy, mercapto, aryl, mono- or di(C<sub>1</sub>-4alkyl) amino, C<sub>1</sub>-4 alkyloxy and pyridinyl;

20 C<sub>1</sub>-4alkyloxycarbonyl; aryl; C<sub>1</sub>-4alkylcarbonyl; C<sub>1</sub>-4alkylthiocarbonyl; arylcarbonyl; arylthiocarbonyl; arylaminocarbonyl; arylaminothiocarbonyl; C<sub>3</sub>-7cycloalkyl;

25 C<sub>1</sub>-4alkane-diyl-C(=O)-Z-C<sub>1</sub>-6alkyl; -C(=O)-Z-C<sub>1</sub>-6alkyl;

-Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-C<sub>1-6</sub>alkyl and R<sup>6</sup>;

aryl represents phenyl optionally substituted with one, two or three substituents each independently selected from nitro, azido, cyano, halo, hydroxy, mercapto, C<sub>1-4</sub>alkyl, C<sub>3-7</sub>cycloalkyl, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylthio, formyl, polyhaloC<sub>1-4</sub>alkyl,

5 NR<sup>9</sup>R<sup>10</sup>, C(=O)NR<sup>9</sup>R<sup>10</sup>, C(=O)-Z-R<sup>14</sup>, R<sup>6</sup>, -O-R<sup>6</sup>, phenyl, Het<sup>3</sup>, C(=O)Het<sup>3</sup> and C<sub>1-4</sub>alkyl substituted with one or more substituents each independently selected from halo, hydroxy, mercapto, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylthio, C(=O)-Z-R<sup>14</sup>,

-Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup> or NR<sup>9</sup>R<sup>10</sup>;

Het<sup>1</sup> represents a three-membered, four-membered, five-membered or six-

10 membered aromatic or non-aromatic, monocyclic or polycyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus, or a fused polycyclic ring system including such heterocycle (such as for instance a fused benzoheterocycle); non-limiting examples of such heterocycles include for

15 instance pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxanyl, dithianyl,

20 trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each

25 independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het<sup>2</sup>, R<sup>11</sup> and C<sub>1-4</sub>alkyl optionally substituted with one or, where possible, two or three substituents each independently selected from Het<sup>2</sup> and R<sup>11</sup>;

Het<sup>2</sup> represents a three-membered, four-membered, five-membered or six-

30 membered aromatic or non-aromatic, monocyclic or polycyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus, or a fused polycyclic

ring system including such heterocycle (such as for instance a fused benzoheterocycle); non-limiting examples of such heterocycles include for instance pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl,  
5 oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl,  
10 cinnolinyl, phthalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het<sup>4</sup>, R<sup>11</sup> and C<sub>1-4</sub>alkyl optionally substituted with one or, where possible, two or three substituents  
15 each independently selected from Het<sup>4</sup> and R<sup>11</sup>;  
Het<sup>3</sup> represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic monocyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus; non-limiting examples of such heterocycles  
20 include for instance pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxolanyl and tetrahydropyranyl; wherein said monocyclic heterocycles each independently may optionally be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylcarbonyl, piperidinyl, NR<sup>12</sup>R<sup>13</sup>,  
25 C(=O)-Z-R<sup>14</sup>, R<sup>6</sup> and C<sub>1-4</sub>alkyl substituted with one or two substituents independently selected from hydroxy, carbonyl C<sub>1-4</sub>alkyloxy, phenyl, C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, R<sup>6</sup> and NR<sup>12</sup>R<sup>13</sup>;  
Het<sup>4</sup> represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic monocyclic heterocycle comprising one or  
30 more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus; non-limiting examples of such heterocycles include for instance pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl,

thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and triazinyl;

Het<sup>5</sup> represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic, monocyclic or polycyclic heterocycle

5 comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus, or a fused polycyclic ring system including such heterocycle (such as for instance a fused benzoheterocycle); non-limiting examples of such heterocycles include for

instance pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl,

10 triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl,

oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl,

oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl,

piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, tetrahydropyranyl,

dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl,

15 benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl,

indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl,

benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl,

quinoxaliny, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl;

wherein said heterocycles each independently may be substituted with, where

20 possible, one, two, three or four substituents each independently selected from

hydroxy, mercapto, carbonyl, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylthio,

C<sub>1-4</sub>alkylcarbonyl, piperidinyl, NR<sup>17</sup>R<sup>18</sup>, C(=O)-Z-C<sub>1-6</sub>alkyl, R<sup>6</sup>, sulfonamido and

C<sub>1-4</sub>alkyl substituted with one or two substituents independently selected from

hydroxy, C<sub>1-4</sub>alkyloxy, mercapto, C<sub>1-4</sub>alkylthio, phenyl, C(=O)-Z-C<sub>1-6</sub>alkyl,

25 -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-C<sub>1-6</sub>alkyl, R<sup>6</sup> and NR<sup>17</sup>R<sup>18</sup>;

Het<sup>6</sup> represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic monocyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus; non-limiting examples of such heterocycles

30 include for instance pyrrolidinyl, piperidinyl, azaridinyl, pyrazolinyl and pyrolinyl,

wherein said heterocycle may optionally be substituted with one, or where

possible, two or three substituents each independently selected from Het<sup>2</sup>, R<sup>11</sup>

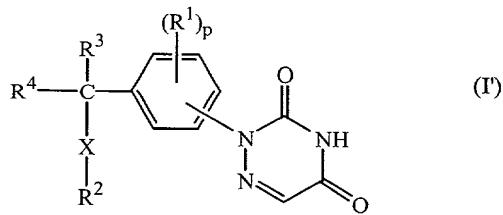
and C<sub>1-4</sub>alkyl optionally substituted with one or more substituents independently selected from Het<sup>2</sup> and R<sup>11</sup>.

provided however that

- R<sup>2</sup> is other than C<sub>1-6</sub> alkyloxycarbonylC<sub>1-6</sub>alkyl or aminocarbonyl; and
- 5 • R<sup>7</sup>, R<sup>8</sup>, R<sup>9</sup> and R<sup>10</sup> are other than aminocarbonyl, C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxyC<sub>1-4</sub>alkylcarbonyl, C<sub>1-4</sub>alkyloxycarbonylcarbonyl, C(=O)-O-R<sup>19</sup>, C<sub>1-4</sub>alkanediyilC(=O)-O-R<sup>19</sup> or -Y-C<sub>1-4</sub>alkanediyilC(=O)-O-R<sup>19</sup>; and
- 10 • R<sup>12</sup> and R<sup>13</sup> are other than C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxyC<sub>1-4</sub>alkylcarbonyl or C<sub>1-4</sub>alkylcarbonylcarbonyl; and
- R<sup>11</sup> is other than C(=O)-O-R<sup>19</sup>, Y-C<sub>1-4</sub>alkanediyil – C(=O)-OR<sup>19</sup>, C(=O)NH<sub>2</sub>, C(=O)NHC<sub>1-4</sub>alkyl or C(=O)NHC<sub>3-7</sub>cycloalkyl; and
- 15 • R<sup>15</sup> and R<sup>16</sup> are other than aminocarbonyl, C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl or C<sub>1-4</sub>alkyloxycarbonylcarbonyl; and
- aryl is other than phenyl substituted with C(=O)-O-R<sup>19</sup>, C(=O)NH<sub>2</sub>, C(=O)NHC<sub>1-4</sub>alkyl, C(=O)NHC<sub>3-7</sub>cycloalkyl and/or with C<sub>1-4</sub>alkyl substituted with C(=O)-O-R<sup>19</sup> or Y-C<sub>1-4</sub>alkanediyil – C(=O)-O-R<sup>14</sup>; and
- 20 • Het<sup>3</sup> is other than a monocyclic heterocycle substituted with C(=O)-O-R<sup>19</sup> and/or with C<sub>1-4</sub>alkyl substituted with C(=O)-O-R<sup>19</sup> and/or Y-C<sub>1-4</sub>alkanediyil C(=O)-O-R<sup>19</sup>; and
- in each of the above proviso's R<sup>19</sup> is defined as hydrogen, C<sub>1-4</sub>alkyl, C<sub>3-7</sub>cycloalkyl, aminocarbonylmethylene or mono- or di(C<sub>1-4</sub>alkyl)aminocarbonylmethylene; and

25 wherein the said compound having the formula (I) contains at least one -C(=O)-Z-R<sup>14</sup> moiety.

2. A compound according to claim 1 having the formula



a *N*-oxide, a pharmaceutically acceptable addition salt or a stereochemically isomeric form thereof, wherein :

*p* represents an integer being 0, 1, 2, 3 or 4;

5    *X* represents O, S, NR<sup>5</sup> or a direct bond or *X*-R<sup>2</sup> taken together may represent cyano;

*Y* represents O, S, NR<sup>5</sup>, or S(O)<sub>2</sub>;

each R<sup>1</sup> independently represents C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkyl, halo, polyhaloC<sub>1-6</sub>alkyl, hydroxy, mercapto, C<sub>1-6</sub>alkyloxy, C<sub>1-6</sub>alkylthio, C<sub>1-6</sub>alkylcarbonyloxy,

10    aryl, cyano, nitro, Het<sup>3</sup>, R<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup> or C<sub>1-4</sub>alkyl substituted with C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, R<sup>6</sup> or NR<sup>7</sup>R<sup>8</sup>;

R<sup>2</sup> represents Het<sup>1</sup>, C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkyl substituted with one or two substituents selected from C(=O)-Z-R<sup>14</sup>, hydroxy, cyano, amino, mono- or di(C<sub>1-4</sub>alkyl)amino,

15    C<sub>1-6</sub>alkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkylsulfonyloxy, C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, aryl, aryloxy, arylthio, Het<sup>1</sup>, Het<sup>1</sup>oxy and Het<sup>1</sup>thio; and if X is O, S or NR<sup>5</sup>, then R<sup>2</sup> may also represent aminothiocarbonyl, C<sub>1-4</sub>alkylcarbonyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-4</sub>alkylthiocarbonyl optionally substituted with C(=O)-Z-R<sup>14</sup>, ,

20    arylcarbonyl, arylthiocarbonyl, Het<sup>1</sup>carbonyl or Het<sup>1</sup>thiocarbonyl;

R<sup>3</sup> represents hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl;

R<sup>4</sup> represents hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl; or

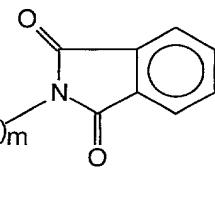
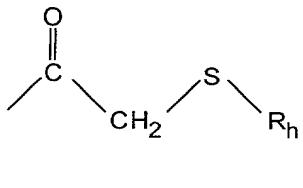
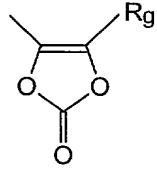
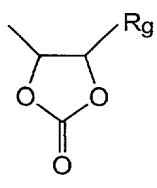
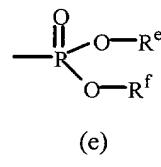
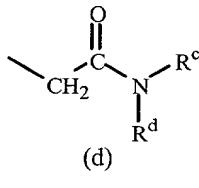
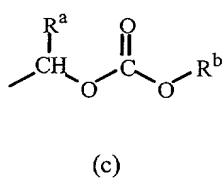
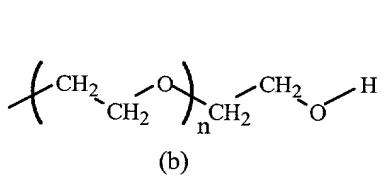
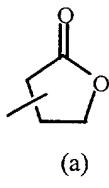
R<sup>3</sup> and R<sup>4</sup> taken together form a C<sub>2-6</sub>alkanediyl;

R<sup>5</sup> represents hydrogen or C<sub>1-4</sub>alkyl;

25    each R<sup>6</sup> independently represents C<sub>1-6</sub>alkylsulfonyl, aminosulfonyl, mono- or di-(C<sub>1-4</sub>alkyl)aminosulfonyl, mono- or di(benzyl)aminosulfonyl,

polyhaloC<sub>1</sub>-6alkylsulfonyl, C<sub>1</sub>-6alkylsulfinyl, phenylC<sub>1-4</sub>alkylsulfonyl, piperazinylsulfonyl, aminopiperidinylsulfonyl, piperidinylaminosulfonyl, N-C<sub>1-4</sub>alkyl-N-piperidinylaminosulfonyl or mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkylsulfonyl; each R<sup>7</sup> and each R<sup>8</sup> are independently selected from hydrogen, C<sub>1-4</sub>alkyl,  
5 hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, aryl, arylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, arylcarbonyl, Het<sup>3</sup>carbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het<sup>3</sup>aminocarbonyl, Het<sup>3</sup>aminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, Het<sup>4</sup> and R<sup>6</sup>;  
10 R<sup>9</sup> and R<sup>10</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl, hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl, phenylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, phenylcarbonyl, Het<sup>3</sup>carbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, Het<sup>3</sup>aminocarbonyl, Het<sup>3</sup>aminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, Het<sup>4</sup> and R<sup>6</sup>;  
15 each R<sup>11</sup> independently being selected from hydroxy, mercapto, cyano, nitro, halo, trihalomethyl, C<sub>1-4</sub>alkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>, formyl, trihaloC<sub>1-4</sub>alkylsulfonyloxy, R<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup>, C(=O)NR<sup>15</sup>R<sup>16</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, aryl, aryloxy, arylcarbonyl, C<sub>3-7</sub>cycloalkyl  
20 optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>3-7</sub>cycloalkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>, phthalimide-2-yl, Het<sup>3</sup>, Het<sup>4</sup> and C(=O)Het<sup>3</sup>; R<sup>12</sup> and R<sup>13</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl, hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl, phenylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, phenylcarbonyl, mono- or  
25 di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup> and R<sup>6</sup>;  
each R<sup>14</sup> independently represents C<sub>1-4</sub> alkyl substituted with one or more substituents selected from phenyl, di- C<sub>1-4</sub>alkylamino, cyano, Het<sup>1</sup> and C<sub>3-7</sub> cycloalkyl, hydrogen, C<sub>1-20</sub>acyl (having a straight or branched, saturated or

unsaturated hydrocarbon chain having 1 to 20 carbon atoms), C<sub>1-20</sub>alkyl, C<sub>3-7</sub>cycloalkyl, polyhaloC<sub>1-20</sub>alkyl or a radical of formula



5 wherein n is 0 to 5 and m is 1 to 4;

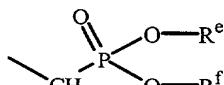
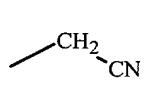
R<sup>a</sup>, R<sup>b</sup>, R<sup>c</sup>, R<sup>d</sup>, R<sup>e</sup> and R<sup>f</sup> are each independently hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl; or

R<sup>e</sup> and R<sup>f</sup> taken together may form -CH<sub>2</sub>-CH<sub>2</sub>-, -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>- or -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-;

10 R<sub>g</sub> and R<sub>h</sub> are each independently C<sub>1-4</sub> alkyl;

each Z independently represents O, S, NH, -CH<sub>2</sub>-O- or -CH<sub>2</sub>-S- whereby -CH<sub>2</sub>- is attached to the carbonyl group;

-Z-R<sup>14</sup> taken together form a radical of formula



(f)

(g)

15 R<sup>15</sup> and R<sup>16</sup> are each independently selected from dihydroxyC<sub>1-4</sub>alkyl, aryl,

arylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, -C(=O)-Z-R<sup>14</sup>, arylcarbonyl, mono- or

di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, arylaminocarbonyl, arylaminothiocarbonyl,

Het<sup>3</sup>aminocarbonyl, Het<sup>3</sup>aminothiocarbonyl, pyridinyLC<sub>1-4</sub>alkyl, Het<sup>3</sup>, Het<sup>4</sup> or R<sup>6</sup>;

aminocarbonylmethylene or mono-or di( $C_{1-4}$ alkyl)aminocarbonylmethylene;  
aryl represents phenyl optionally substituted with one, two or three substituents  
each independently selected from nitro, azido, cyano, halo, hydroxy,  
 $C_{1-4}$ alkyl,  $C_{3-7}$ cycloalkyl,  $C_{1-4}$ alkyloxy, formyl, polyhalo $C_{1-4}$ alkyl,  $NR^9R^{10}$ ,  
5  $C(=O)NR^9R^{10}$ ,  $C(=O)-Z-R^{14}$ ,  $R^6$ , -O- $R^6$ , phenyl,  $Het^3$ ,  $C(=O)Het^3$  and  $C_{1-4}$ alkyl  
substituted with one or more substituents each independently selected from  
halo, hydroxy,  $C_{1-4}$ alkyloxy,  $C(=O)-Z-R^{14}$ , -Y- $C_{1-4}$ alkanediyl-C(=O)-Z-R<sup>14</sup>,  $Het^3$   
or  $NR^9R^{10}$ ;

10  $Het^1$  represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl,  
imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl,  
tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl,  
thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl,  
pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl,  
thiomorpholinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl,  
15 isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl,  
indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl,  
benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl,  
quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl;  
wherein said heterocycles each independently may optionally be substituted  
20 with one, or where possible, two or three substituents each independently  
selected from  $Het^2$ ,  $R^{11}$  and  $C_{1-4}$ alkyl optionally substituted with one or two  
substituents independently selected from  $Het^2$  and  $R^{11}$ ;

25  $Het^2$  represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl,  
imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl,  
tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl,  
thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl,  
pyrazinyl, pyranyl, pyridazinyl, dioxanyl, dithianyl, trithianyl, triazinyl,  
benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl,  
benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-  
30 d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl,  
quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl;  
wherein said heterocycles each independently may optionally be

substituted with one, or where possible, two or three substituents each independently selected from Het<sup>4</sup>, R<sup>11</sup> and C<sub>1-4</sub>alkyl optionally substituted with one or two substituents independently selected from Het<sup>4</sup> and R11;

Het<sup>3</sup> represents a monocyclic heterocycle selected from pyrrolidinyl, piperidinyl,  
5 piperazinyl, morpholinyl, thiomorpholinyl and tetrahydropyran; wherein said monocyclic heterocycles each independently may optionally be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylcarbonyl, piperidinyl, NR<sup>12</sup>R<sup>13</sup>, C(=O)-Z-R<sup>14</sup>, R<sup>6</sup> and C<sub>1-4</sub>alkyl substituted with one or two substituents  
10 independently selected from hydroxy, C<sub>1-4</sub>alkyloxy, phenyl, C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, R<sup>6</sup> and NR<sup>12</sup>R<sup>13</sup>;

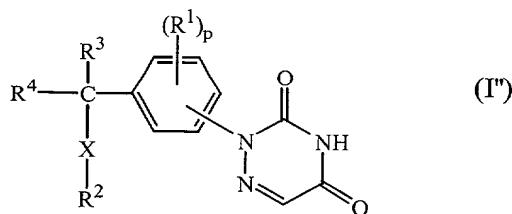
Het<sup>4</sup> represents a monocyclic heterocycle selected from pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl, thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, 15 pyridazinyl and triazinyl provided however that

- R<sup>2</sup> is other than C<sub>1-6</sub> alkyloxycarbonylC<sub>1-6</sub>alkyl, aminocarbonyl; and
- R<sup>7</sup>, R<sup>8</sup>, R<sup>9</sup> and R<sup>10</sup> are other than aminocarbonyl, C<sub>1-4</sub>alkylcarbonyloxy- C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl, C<sub>1-4</sub>alkyloxycarbonylcarbonyl C(=O)-O-R<sup>14</sup>, C<sub>1-4</sub>alkanediylC(=O)-O-R<sup>14</sup> and -Y-C<sub>1-4</sub>alkanediylC(=O)-O-R<sup>14</sup>; and
- R<sup>12</sup> and R<sup>13</sup> are other than C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl, C<sub>1-4</sub>alkylcarbonylcabonyl; and
- R<sup>11</sup> is other than C(=O)-O-R<sup>14</sup>, Y-C<sub>1-4</sub>alkanediyl – C(=O)-OR<sup>14</sup>, C(=O)NH<sub>2</sub>, C(=O)NHC<sub>1-4</sub>alkyl or C(=O)NHC<sub>3-7</sub>cycloalkyl; and
- R<sup>14</sup> is other than hydrogen, C<sub>1-4</sub>alkyl, C<sub>3-7</sub>cycloalkyl, aminocarbonylmethylene, mono- or di (C<sub>1-4</sub>alkyl) aminocarbonylmethylene in the event Z is 0; and
- R<sup>15</sup> and R<sup>16</sup> are other than aminocarbonyl, C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl or C<sub>1-4</sub>alkyloxycarbonylcarbonyl; and
- Aryl is other than phenyl substituted with C(=O)-O-R<sup>14</sup> C(=O)NH<sub>2</sub>, C(=O)NHC<sub>1-4</sub>alkyl, C(=O)NHC<sub>3-7</sub>cycloalkyl and/or with C<sub>1-4</sub>alkyl substituted with C(=O)-O-R<sup>14</sup> or Y-C<sub>1-4</sub>alkanediyl – C(=O)-O-R<sup>14</sup>; and

- Het<sup>3</sup> is other than a monocyclic heterocycle substituted with C(=O)-O-R<sup>14</sup> and/or with C<sub>1-4</sub>alkyl substituted with C(=O)-O-R<sup>14</sup> and/or Y-C<sub>1-4</sub>alkanediyl-(=O)-O-R<sup>14</sup>; and
- The said compound of formula (I) contains at least one -C(=O)-Z-R<sup>14</sup> moiety.

5

3. A compound according to claims 1 or 2 having the formula



a N-oxide, a pharmaceutically acceptable addition salt or a stereochemically isomeric form thereof, wherein :

10 p represents an integer being 0, 1, 2, 3 or 4;  
 X represents O, S, NR<sup>5</sup> or a direct bond or-X-R<sup>2</sup> taken together may represent cyano;

Y represents O, S, NR<sup>5</sup>, or S(O)<sub>2</sub>;

each R<sup>1</sup> independently represents C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkyl, halo, polyhaloC<sub>1-6</sub>alkyl, hydroxy, mercapto, C<sub>1-6</sub>alkyloxy, C<sub>1-6</sub>alkylthio, C<sub>1-6</sub>alkylcarbonyloxy, aryl, cyano, nitro, Het<sup>3</sup>, R<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup> or C<sub>1-4</sub>alkyl substituted with C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, R<sup>6</sup> or NR<sup>7</sup>R<sup>8</sup>;

15 R<sup>2</sup> represents Het<sup>1</sup>, C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkyl substituted with one or two substituents selected from C(=O)-Z-R<sup>14</sup>, hydroxy, cyano, amino, mono- or di(C<sub>1-4</sub>alkyl)amino, C<sub>1-6</sub>alkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-6</sub>alkylsulfonyloxy, C<sub>3-7</sub>cycloalkyl optionally substituted with C(=O)-Z-R<sup>14</sup>, aryl, aryloxy, arylthio, Het<sup>1</sup>, Het<sup>1</sup>oxy and Het<sup>1</sup>thio; and if X is O, S or NR<sup>5</sup>, then R<sup>2</sup> may also represent aminothiocarbonyl, C<sub>1-4</sub>alkylcarbonyl optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>1-4</sub>alkylthiocarbonyl optionally substituted with C(=O)-Z-R<sup>14</sup>, arylcarbonyl, arylthiocarbonyl, Het<sup>1</sup>carbonyl or Het<sup>1</sup>thiocarbonyl;

20 R<sup>3</sup> represents hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl;

25 R<sup>4</sup> represents hydrogen, C<sub>1-6</sub>alkyl or C<sub>3-7</sub>cycloalkyl; or

R<sup>3</sup> and R<sup>4</sup> taken together form a C<sub>2-6</sub>alkanediyl;

R<sup>5</sup> represents hydrogen or C<sub>1-4</sub>alkyl;

each R<sup>6</sup> independently represents C<sub>1-6</sub>alkylsulfonyl, aminosulfonyl,

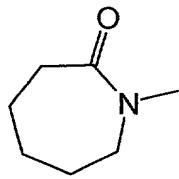
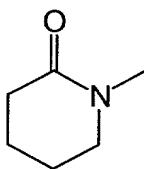
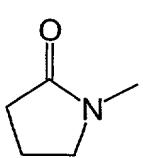
piperidinylsulfonyl, mono- or di(C<sub>1-4</sub>alkyl)aminosulfonyl, mono- or

5 di(benzyl)aminosulfonyl, polyhaloC<sub>1-6</sub>alkylsulfonyl, C<sub>1-6</sub>alkylsulfinyl, phenylC<sub>1-4</sub>alkylsulfonyl, piperazinylsulfonyl, aminopiperidinylsulfonyl, piperidinylaminosulfonyl,

N-C<sub>1-4</sub>alkyl-N-piperidinylaminosulfonyl or mono-or

di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkylsulfonyl;

10 each R<sup>7</sup> and each R<sup>8</sup> are independently selected from hydrogen, C<sub>1-4</sub>alkyl, hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, aryl, arylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, arylcarbonyl, Het<sup>3</sup>carbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het<sup>3</sup>aminocarbonyl, Het<sup>3</sup>aminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, Het<sup>4</sup> and R<sup>6</sup>; or R<sup>7</sup> and R<sup>8</sup> taken together with the nitrogen atom to which they are attached form a radical of formula



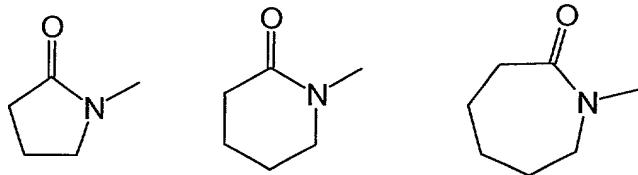
20

R<sup>9</sup> and R<sup>10</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl,

hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl, phenylC<sub>1-4</sub>alkyl,

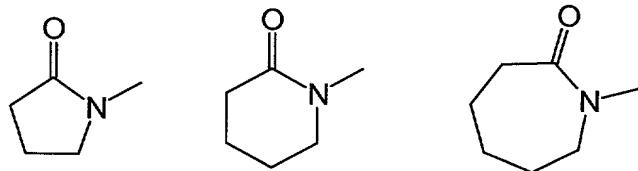
C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, phenylcarbonyl, Het<sup>3</sup>carbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, phenylaminocarbonyl,

25 phenylaminothiocarbonyl, Het<sup>3</sup>aminocarbonyl, Het<sup>3</sup>aminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup>, Het<sup>4</sup> and R<sup>6</sup>; or R<sup>9</sup> and R<sup>10</sup> taken together with the nitrogen atom to which they are attached form a radical of formula



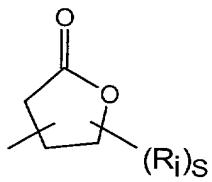
each R<sup>11</sup> independently being selected from hydroxy, mercapto, cyano, nitro,  
5 halo, trihalomethyl, C<sub>1-4</sub>alkyloxy optionally substituted with C(=O)-Z-R<sup>14</sup>,  
formyl, trihaloC<sub>1-4</sub>alkylsulfonyloxy, R<sup>6</sup>, NR<sup>7</sup>R<sup>8</sup>, C(=O)NR<sup>15</sup>R<sup>16</sup>, -C(=O)-Z-R<sup>14</sup>, -  
Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, aryl, aryloxy, arylcarbonyl, C<sub>3-7</sub>cycloalkyl  
optionally substituted with C(=O)-Z-R<sup>14</sup>, C<sub>3-7</sub>cycloalkyloxy optionally  
substituted with C(=O)-Z-R<sup>14</sup>, phthalimide-2-yl, Het<sup>3</sup>, Het<sup>4</sup> and C(=O)Het<sup>3</sup>;

10 R<sup>12</sup> and R<sup>13</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl,  
hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl, phenylC<sub>1-4</sub>alkyl,  
C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, phenylcarbonyl, mono- or  
di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl,  
C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, -C(=O)-Z-R<sup>14</sup>, -  
Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup> and R<sup>6</sup>; or R<sup>12</sup> and R<sup>13</sup> taken together with the  
15 nitrogen atom to which they are attached form a radical of formula

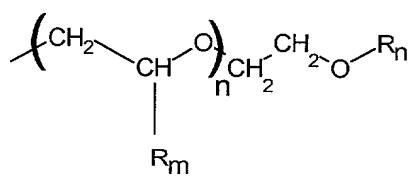


20 each R<sup>14</sup> independently represents hydrogen, C<sub>1-20</sub>acyl (having a straight or  
branched, saturated or unsaturated hydrocarbon chain having 1 to 20  
carbon atoms), C<sub>1-20</sub>alkyl, C<sub>3-20</sub>alkenyl optionally substituted with phenyl,  
C<sub>3-20</sub>alkynyl, C<sub>3-7</sub> cycloalkyl, polyhaloC<sub>1-20</sub>alkyl, Het<sup>5</sup>, phenyl or C<sub>1-20</sub> alkyl  
substituted with one or more substituents selected from hydroxy, NR<sup>17</sup>R<sup>18</sup>,  
25 phenyl, mono- or di-(C<sub>1-4</sub>alkyl)amino, cyano, Het<sup>5</sup>, C<sub>1-4</sub> alkyloxycarbonyl,

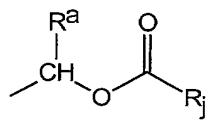
phenyl C<sub>1-4</sub> alkyloxycarbonyl and C<sub>3-7</sub> cycloalkyl, or R<sup>14</sup> represents a radical of formula



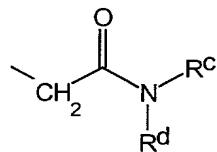
(a)



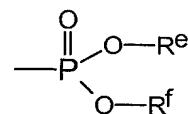
(b)



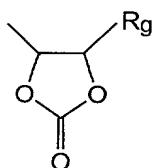
(c)



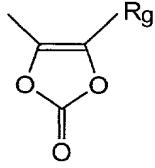
(d)



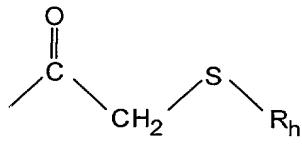
(e)



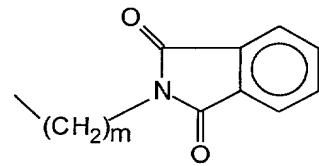
(h)



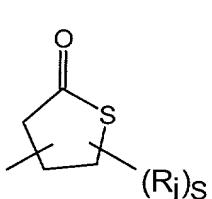
(i)



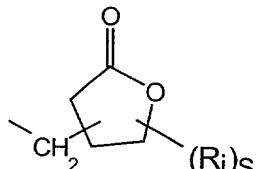
(j)



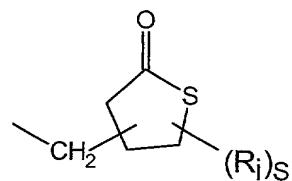
(k)



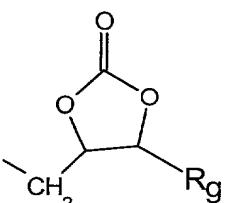
(I)



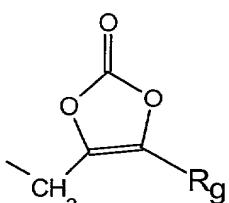
(m)



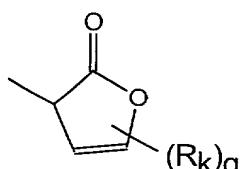
(n)



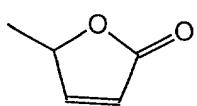
(o)



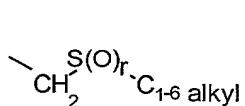
(p)



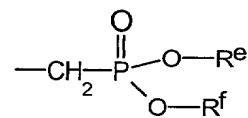
(q)



(r)



(s)



(t)

wherein m is 1 to 4, n is 0 to 5, q is 0 to 2, r is 0 to 2 and s is 0 to 4;

R<sup>a</sup>, R<sup>b</sup>, R<sup>c</sup>, R<sup>d</sup>, R<sup>e</sup> and R<sup>f</sup> are each independently hydrogen, C<sub>1-6</sub>alkyl, phenyl or

5 C<sub>3-7</sub>cycloalkyl; or

R<sup>e</sup> and R<sup>f</sup> taken together may form -CH<sub>2</sub>-CH<sub>2</sub>-, -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>- or -CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-;

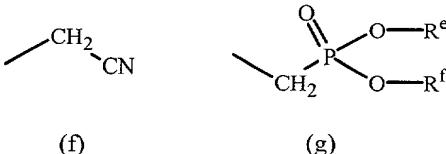
R<sub>g</sub>, R<sub>h</sub> and R<sub>k</sub> are each independently hydrogen or C<sub>1-4</sub> alkyl;

R<sub>i</sub> is C<sub>1-4</sub>alkyl;

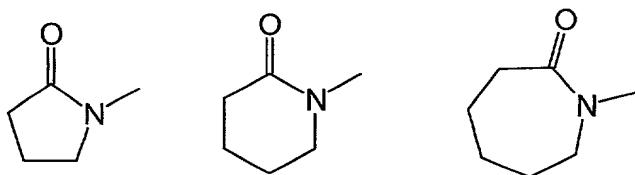
10 R<sub>j</sub> is -O-R<sub>b</sub>, C<sub>1-6</sub>alkyl, phenyl or C<sub>3-7</sub>cycloalkyl optionally substituted with C<sub>1-4</sub> alkyloxy;

where R<sub>m</sub> is hydrogen or C<sub>1-4</sub> alkyloxy and R<sub>n</sub> is hydrogen, C<sub>1-4</sub>alkyl, C<sub>3-7</sub>cycloalkyl, phenyl or phenylC<sub>1-4</sub>alkyl

each Z independently represents O, S, NH, -CH<sub>2</sub>-O- or -CH<sub>2</sub>-S- whereby -CH<sub>2</sub>- is attached to the carbonyl group; or  
-Z-R<sup>14</sup> taken together form a radical of formula



5      R<sup>15</sup> and R<sup>16</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl,  
hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, aryl, aryIC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl,  
-C(=O)-Z-R<sup>14</sup>, arylcarbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl,  
arylaminoacarbonyl, arylaminothiocarbonyl, aminocarbonylmethylene, mono-  
or di(C<sub>1-4</sub>alkyl) aminocarbonylmethylene, Het<sup>3</sup>aminocarbonyl,  
10     Het<sup>3</sup>aminothiocarbonyl,  
pyridinylC<sub>1-4</sub>alkyl, Het<sup>3</sup> or R<sup>6</sup>; or R<sup>15</sup> and R<sup>16</sup> taken together with the nitrogen  
atom to which they are attached form a radical of formula



15 R<sup>17</sup> and R<sup>18</sup> are each independently selected from hydrogen, C<sub>1-4</sub>alkyl, hydroxyC<sub>1-4</sub>alkyl, dihydroxyC<sub>1-4</sub>alkyl, phenyl, phenylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxyC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkylcarbonyl, phenylcarbonyl, mono- or di(C<sub>1-4</sub>alkyl)aminoC<sub>1-4</sub>alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, C<sub>3-7</sub>cycloalkyl, pyridinylC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanediyl-C(=O)-Z-C<sub>1-6</sub>alkyl, -C(=O)-Z-C<sub>1-6</sub>alkyl, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-C<sub>1-6</sub>alkyl and R<sup>6</sup>;  
20 aryl represents phenyl optionally substituted with one, two or three substituents each independently selected from nitro, azido, cyano, halo, hydroxy, C<sub>1-4</sub>alkyl, C<sub>3-7</sub>cycloalkyl, C<sub>1-4</sub>alkyloxy, formyl, polyhaloC<sub>1-4</sub>alkyl, NR<sup>9</sup>R<sup>10</sup>, C(=O)NR<sup>9</sup>R<sup>10</sup>, C(=O)-Z-R<sup>14</sup>, R<sup>6</sup>, -O-R<sup>6</sup>, phenyl, Het<sup>3</sup>, C(=O)Het<sup>3</sup> and C<sub>1-4</sub>alkyl  
25 substituted with one or more substituents each independently selected from

halo, hydroxy, C<sub>1-4</sub>alkyloxy, C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, Het<sup>3</sup> or NR<sup>9</sup>R<sup>10</sup>;

Het<sup>1</sup> represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, 5 tetrahydrofuran, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, 10 benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl, isoindolyl, indolinyl, purinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phthalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het<sup>2</sup>, R<sup>11</sup> and 15 C<sub>1-4</sub>alkyl optionally substituted with one or two substituents independently selected from Het<sup>2</sup> and R<sup>11</sup>;

Het<sup>2</sup> represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, 20 tetrahydrofuran, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 25 1H-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phthalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het<sup>4</sup>, R<sup>11</sup> and C<sub>1-4</sub>alkyl optionally substituted with one or two substituents independently selected from Het<sup>4</sup> and R<sup>11</sup>;

Het<sup>3</sup> represents a monocyclic heterocycle selected from pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl and tetrahydropyranyl; wherein said monocyclic heterocycles each independently may optionally be substituted with, where possible, one, two, three or four substituents each independently

5 selected from hydroxy, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylcarbonyl, piperidinyl, NR<sup>12</sup>R<sup>13</sup>, C(=O)-Z-R<sup>14</sup>, R<sup>6</sup> and C<sub>1-4</sub>alkyl substituted with one or two substituents independently selected from hydroxy, C<sub>1-4</sub>alkyloxy, phenyl, C(=O)-Z-R<sup>14</sup>, -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-R<sup>14</sup>, R<sup>6</sup> and NR<sup>12</sup>R<sup>13</sup>;

Het<sup>4</sup> represents a monocyclic heterocycle selected from pyrrolyl, imidazolyl, 10 pyrazolyl, triazolyl, tetrazolyl, furanyl, thieryl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and triazinyl;

Het<sup>5</sup> represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl,

15 tetrahydrofuranlyl, thieryl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl,

pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, tetrahydropyranyl, dioxanyl,

dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl,

20 isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl,

isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl,

quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl,

thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said

25 heterocycles each independently may be substituted with, where possible,

one, two, three or four substituents each independently selected from

hydroxy, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, C<sub>1-4</sub>alkylcarbonyl, piperidinyl, NR<sup>17</sup>R<sup>18</sup>,

C(=O)-Z-C<sub>1-6</sub>alkyl, R<sup>6</sup>, sulfonamido and C<sub>1-4</sub>alkyl substituted with one or two

substituents independently selected from hydroxy, C<sub>1-4</sub>alkyloxy, phenyl,

C(=O)-Z-C<sub>1-6</sub>alkyl,

30 -Y-C<sub>1-4</sub>alkanediyl-C(=O)-Z-C<sub>1-6</sub>alkyl, R<sup>6</sup> and NR<sup>17</sup>R<sup>18</sup>;

provided however that

- R<sup>2</sup> is other than C<sub>1-6</sub> alkyloxycarbonylC<sub>1-6</sub>alkyl or aminocarbonyl; and

- R<sup>7</sup>, R<sup>8</sup>, R<sup>9</sup> and R<sup>10</sup> are other than aminocarbonyl, C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl, C<sub>1-4</sub>alkyloxycarbonylcarbonyl, C(=O)-O-R<sup>19</sup>, C<sub>1-4</sub>alkanediylC(=O)-O-R<sup>19</sup> or -Y-C<sub>1-4</sub>alkanediylC(=O)-O-R<sup>19</sup>; and

5     • R<sup>12</sup> and R<sup>13</sup> are other than C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl or C<sub>1-4</sub>alkylcarbonylcarbonyl; and

- R<sup>11</sup> is other than C(=O)-O-R<sup>19</sup>, Y-C<sub>1-4</sub>alkanediyl – C(=O)-OR<sup>19</sup>, C(=O)NH<sub>2</sub>, C(=O)NHC<sub>1-4</sub>alkyl or C(=O)NHC<sub>3-7</sub>cycloalkyl; and

- R<sup>15</sup> and R<sup>16</sup> are other than aminocarbonyl, C<sub>1-4</sub>alkylcarbonyloxy-C<sub>1-4</sub>alkylcarbonyl, hydroxy C<sub>1-4</sub>alkylcarbonyl or C<sub>1-4</sub>alkyloxycarbonylcarbonyl; and

- aryl is other than phenyl substituted with C(=O)-O-R<sup>19</sup>, C(=O)NH<sub>2</sub>, C(=O)NHC<sub>1-4</sub>alkyl, C(=O)NHC<sub>3-7</sub>cycloalkyl and/or with C<sub>1-4</sub>alkyl substituted with C(=O)-O-R<sup>19</sup> or Y-C<sub>1-4</sub>alkanediyl – C(=O)-O-R<sup>14</sup>; and

- Het<sup>3</sup> is other than a monocyclic heterocycle substituted with C(=O)-O-R<sup>19</sup> and/or with C<sub>1-4</sub>alkyl substituted with C(=O)-O-R<sup>19</sup> and/or Y-C<sub>1-4</sub>alkanediyl – (=O)-O-R<sup>19</sup>; and

- in each of the above proviso's R<sup>19</sup> is defined as hydrogen, C<sub>1-4</sub>alkyl, C<sub>3-7</sub>cycloalkyl, aminocarbonylmethylene or mono- or di(C<sub>1-4</sub>alkyl)aminocarbonylmethylene; and

- the said compound of formula (I) contains at least one – C(=O)-Z-R<sup>14</sup> moiety.

4. A compound according to any of claims 1 to 3 wherein the 6-azauracil

25     moiety is in the para position relative to the carbon atom bearing the -X-R<sup>2</sup>, R<sup>3</sup> and R<sup>4</sup> substituents.

5. A compound according to any of claims 1 to 4 wherein R<sup>2</sup> is a monocyclic heterocycle selected from pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl, thieryl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and

triazinyl, wherein said monocyclic heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het<sup>2</sup>, R<sup>11</sup> and C<sub>1-4</sub>alkyl optionally substituted with Het<sup>2</sup> or R<sup>11</sup>.

5

6. A compound according to any of claims 1 to 5 wherein R<sup>3</sup> and R<sup>4</sup> are both methyl and -X-R<sup>2</sup> is Het<sup>1</sup>.

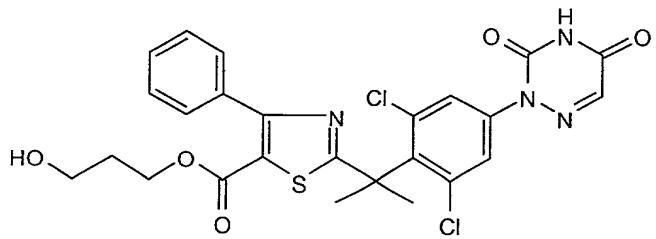
10 7. A compound according to any of claims 1 to 6 wherein p is 1 or 2 and each R<sup>1</sup> is chloro.

15 8. A compound according to any of claims 1 to 7 wherein R<sup>3</sup> and R<sup>4</sup> are both methyl, -X-R<sup>2</sup> is optionally substituted 2-thiazolyl or 3-oxadiazolyl, the 6-azauracil moiety is in the para position relative to the carbon atom bearing the -X-R<sup>2</sup>, R<sup>3</sup> and R<sup>4</sup> substituents, and p is 2 whereby both R<sup>1</sup> substituents are chloro positioned ortho relative to the carbon atom bearing the -X-R<sup>2</sup>, R<sup>3</sup> and R<sup>4</sup> substituents.

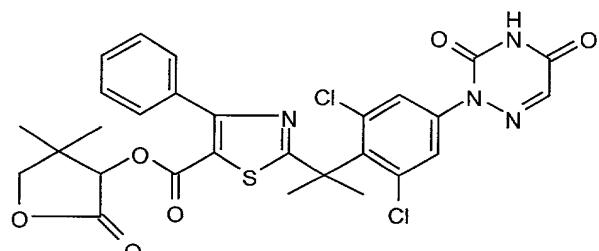
20 9. A compound according to claim 8 wherein X-R<sup>2</sup> is di-substituted with phenyl and either (i) R<sup>11</sup> where R<sup>11</sup> is a group of formula -C(=O)-Z-R<sup>14</sup> in which Z is O and R<sup>14</sup> is C<sub>1-20</sub>alkyl substituted with hydroxy or with Het<sup>5</sup> where Het<sup>5</sup> is piperazinyl substituted with Het<sup>6</sup>sulfonyl, or R<sup>14</sup> is a radical of formula (a) in which R<sub>j</sub> is C<sub>1-6</sub>alkyl and s is 2, or (ii) C<sub>1-4</sub>alkyl substituted with R<sup>11</sup> where R<sup>11</sup> is a group of formula -C(=O)-Z-R<sup>14</sup> in which Z is O and R<sup>14</sup> is a radical of formula (a) in which R<sub>j</sub> is C<sub>1-6</sub>alkyl and s is 2.

25 10 A compound according to claim 1 selected from those of formulae (A), (B), (C) and (D) below:-:

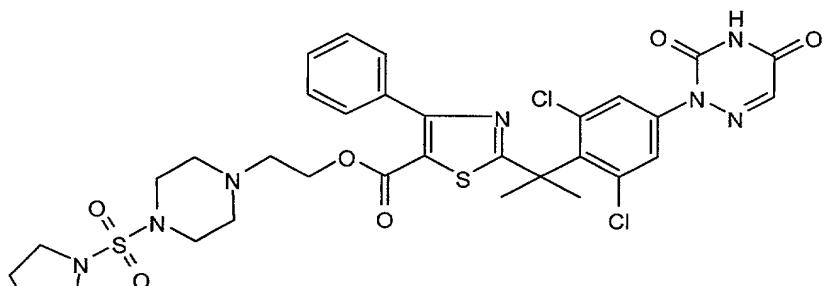
2004TE000020052007



(A)

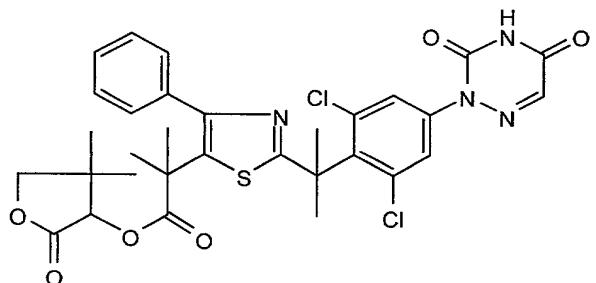


(B)



5

(C)



(D)

11. A composition comprising a pharmaceutically acceptable carrier and, as active ingredient, a therapeutically effective amount of a compound according to any of claims 1 to 10.

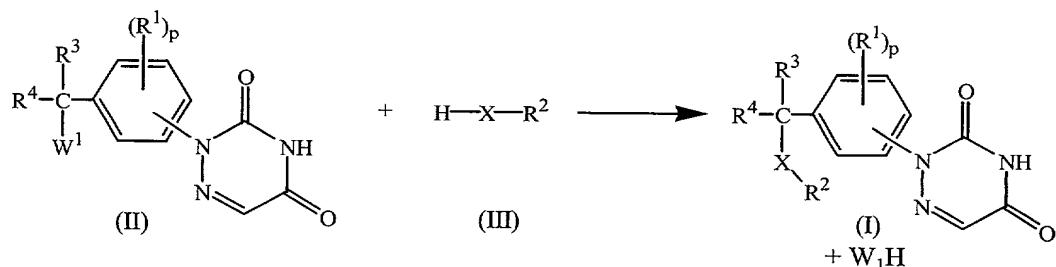
5      12. A process for preparing a composition as claimed in claim 11, wherein a pharmaceutically acceptable carrier is intimately mixed with a therapeutically effective amount of a compound according to any of claims 1 to 10.

10     13. A compound as claimed in any one of claims 1 to 10 for use as a medicine.

14. Use of a compound according to any of claims 1 to 10 in the manufacture of a medicament for treating eosinophil-dependent inflammatory diseases.

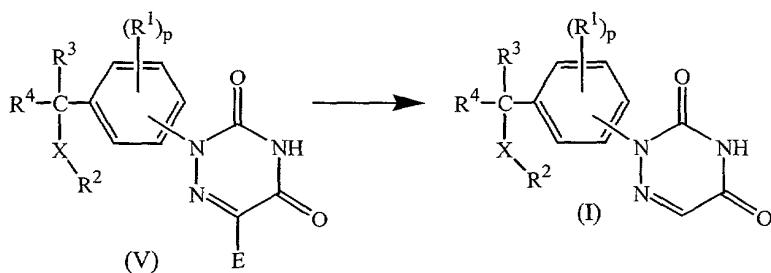
15     15. A process for preparing a compound as claimed in claim 1, comprising the step of

a) reacting an intermediate of formula (II) wherein  $W^1$  is a suitable leaving group with an appropriate reagent of formula (III) optionally in a reaction-inert solvent and optionally in the presence of a base at a temperature ranging between  $-70^\circ\text{C}$  and reflux temperature;



wherein  $R^1$   $R^2$ ,  $R^3$ ,  $R^4$ ,  $p$  and  $X$  are as defined in claim 1 or;

b) eliminating the group E of a triazinedione of formula (V)



wherein E is an appropriate electron attracting group and R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, X and p are as defined in claim 1; and, if desired, converting compounds of formula (I) into each other following art-known transformations, and further, if desired,

- 5     converting the compounds of formula (I), into a therapeutically active non-toxic acid addition salt by treatment with an acid, or into a therapeutically active non-toxic base addition salt by treatment with a base, or conversely, converting the acid addition salt form into the free base by treatment with alkali, or converting the base addition salt into the free acid by treatment with acid; and also, if
- 10    desired, preparing stereochemically isomeric forms or N-oxide forms thereof.

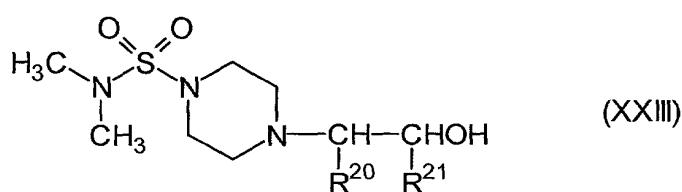
16. A process of marking a receptor comprising the steps of

- a) radiolabelling a compound as defined in claim 1;
- b) administering said radiolabelled compound to biological material,
- 15    c) detecting the emissions from the radiolabelled compound.

17 A process of imaging an organ, characterized by, administering a sufficient amount of a radiolabelled compound of formula (I) in an appropriate composition, and detecting the emissions from the radioactive compound.

20

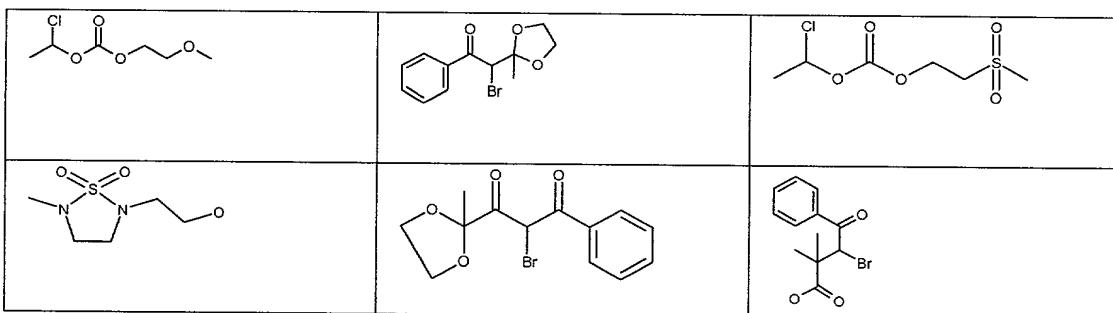
18. A compound of formula



wherein R<sup>20</sup> and R<sup>21</sup> are each independently selected from hydrogen or C<sub>1-20</sub> alkyl or R<sup>20</sup> and R<sup>21</sup> taken together with the carbon atom to which they are attached form a cycloalkyl radical.

5 19. Use of a compound of claim 18 for preparing a compound of claim 1  
wherein Het<sup>5</sup> represents a sulfonamido substituted piperazine.

20. A compound having any of the following formulae:



10 21. Use of a compound of claim 20 as an intermediate for preparing a compound of claim 1.